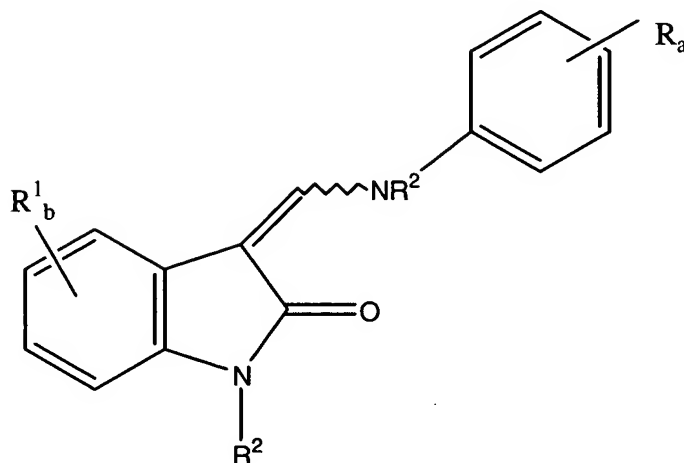


1. A compound represented by the general formula I



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- wherein R^1 is selected from the group consisting of halogen, NO_2 , CN , C_1 to C_4 alkyl and aryl; R^2 is selected from the group consisting of hydrogen, C_1 to C_8 alkyl, COCH_3 , $\text{CH}_2\text{CH}_2\text{OH}$, $\text{CH}_2\text{CH}_2\text{CH}_2\text{OH}$ and phenyl; R is selected from the group consisting of D, halogen, C_1 to C_8 alkyl, CF_3 , OCF_3 , OCF_2H , CH_2CN , CN , SR^2 ,
 10 $(\text{CR}_7\text{R}_8)_c\text{C}(\text{O})\text{OR}^2$, $\text{C}(\text{O})\text{N}(\text{R}^2)_2$, $(\text{CR}_7\text{R}_8)_c\text{OR}^2$, $\text{HNC}(\text{O})\text{R}^2$, $\text{HN}-\text{C}(\text{O})\text{OR}^2$, $(\text{C}\text{R}_7\text{R}_8)_c\text{N}(\text{R}^2)_2$, SO_2 , $(\text{CR}_7\text{R}_8)_c\text{N}(\text{R}^2)_2$, $\text{OP}(\text{O})(\text{OR}^2)_2$, $\text{OC}(\text{O})\text{OR}^2$, OCH_2O , $\text{HN}-\text{CH}=\text{CH}$, $-\text{N}(\text{COR}^2)\text{CH}_2\text{CH}_2$, $\text{HC}=\text{N}-\text{NH}$, $\text{N}=\text{CH}-\text{S}$, $(\text{CR}_7\text{R}_8)_c-\text{R}^6$ and $\text{NR}_2(\text{CR}^7\text{R}^8)_d\text{R}^6$ wherein R^6 is selected from the group consisting of halogen,
 15 dilower alkylamino, piperidinyl, 3-fluoropyrrolidinyl, 3-fluoropiperidinyl, 2-pyridinyl, 3-pyridinyl, 4-pyridinyl, 3-pyrrolinyl, pyrrolidinyl, methyl isonipecotate, $\text{N}-(2\text{-methoxyethyl})-\text{N}-\text{methylamyl}$, 1,2,3,6-tetrahydropyridinyl, morpholinyl, hexamethyleneiminyl, piperazinyl-2-one, piperazinyl, $\text{N}-(2\text{-methoxyethyl})\text{ethylaminyl}$, thiomorpholinyl, heptamethyleneiminyl, 1-
 20 piperazinylcarboxaldehyde, 2,3,6,7-tetrahydro-(1H)-1,4-diazepinyl-5(4H)-one, N-methylhomopiperazinyl, (dimethylamino)pyrrolidinyl, $\text{N}-(2\text{-methoxyethyl})-\text{N}-$

propylaminyl, isoindolinyl, nipecotamidinyl, isonipecotamidinyl, 1-acetylpiperazinyl, 3-acetamidopyrrolidinyl, trans-decahydroisoquinolinyl, cis-decahydroisoquinolinyl, N-acetylhomopiperazinyl, 3-(diethylamino)pyrrolidinyl, 1,4-dioxo-8-azaspiro[4.5]decanyl, 1-(2-methoxyethyl)-piperazinyl, 2-pyrrolidin-3-ylpyridinyl, 4-pyrrolidin-3-ylpyridinyl, 3-(methylsulfonyl)pyrrolidinyl, 3-picolylmethylaminyl, 2-(2-methylaminoethyl)pyridinyl, 1-(2-pyrimidyl)-piperazinyl, 1-(2-pyrazinyl)-piperazinyl, 2-methylaminomethyl-1,3-dioxolane, 2-(N-methyl-2-aminoethyl)-1,3-dioxolane, 3-(N-acetyl-N-methylamino)pyrrolidinyl, 2-methoxyethylaminyl, tetrahydrofurfurylaminyl, 4-aminotetrahydropyran, 2-amino-1-methoxybutane, 2-methoxyisopropylaminyl, 1-(3-aminopropyl)imidazole, histaminyl, N,N-diisopropylethylenediaminyl, 1-benzyl-3-aminopyrrolidyl 2-(aminomethyl)-5-methylpyrazinyl, 2,2-dimethyl-1,3-dioxolane-4-methanaminyl, (R)-3-amino-1-N-BOC-pyrrolidinyl, 4-Amino-1,2,2,6,6-pentamethylpiperidinyl, 4-aminomethyltetrahydropyran, ethanolamine and lower alkyl-substituted derivatives thereof and wherein when c is 1 said CH₂ may be



and CH₂CH₂CH₂; provided said alkyl or phenyl radicals may be substituted with one or two halo, hydroxy or lower alkyl amino radicals and wherein R⁷ and R⁸ are selected from the group consisting of H, F and C₁-C₄ alkyl or CR⁷R⁸ may represent a carbocyclic ring of from 3 to 6 carbons;

b is 0 or an integer of from 1 to 3;

a is 0 or an integer of from 1 to 5;

c is 0 or an integer of from 1 to 4;

d is an integer of from 1 to 5;

the wavy line represents a E or Z bond and pharmaceutically acceptable salts

thereof; provided however said compound is not

- 3-Phenylaminomethylene-1,3-dihydro-indol-2-one,
3-[(3-Bromo-phenylamino)-methylene]-1,3-dihydro-indol-2-one,
3-[(4-Bromo-phenylamino)-methyl]-1,3-dihydro-indol-2-one,
3-[(3-Bromo-phenylamino)-methyl]-1,3-dihydro-indol-2-one,
5 3-[(4-Ethyl-phenylamino)-methylene]-1,3-dihydro-indol-2-one,
3-Ethyl-phenylamino)-methylene]-1,3-dihydro-indol-2-one,
3-[(4-Methoxy-phenylamino)-methylene]-1,3-dihydro-indol-2-one,
4-[(2-Oxo-1,3-dihydro-indol-3-ylididenemethyl)-amino]-benzoic acid ethyl ester,
3-[(2-Ethyl-phenylamino)-methylene]-1,3-dihydro-indol-2-one,
10 3-[(3-Fluoro-4-methyl-phenylamino)-methylene]-1,3-dihydro-indol-2-one,
3-[(3-Fluoro-2-methyl-phenylamino)-methylene]-1,3-dihydro-indol-2-one,
3-[(4-Hydroxy-phenylamino)-methylene]-1,3-dihydro-indol-2-one,
3-[(3-Chloro-4-hydroxy-phenylamino)-methylene]-1,3-dihydro-indol-2-one or
3-[(4-Fluoro-2-methyl-phenylamino)-methylene]-1,3-dihydro-indol-2-one.

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2. The compound of claim 1 wherein R^1 is selected from the group consisting of CH_3 , F and Cl.

3. The compound of claim 1 wherein a is 0.

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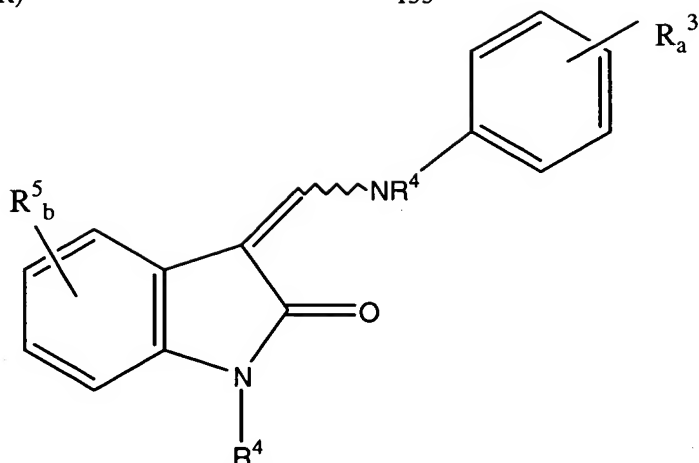
4. The compound of claim 1 wherein R^2 is H.

5. The compound of claim 1 wherein R is selected from the group consisting of CH_3 , CH_2CH_3 , OCH_3 , OH, t-butyl, F, CN, $C(O)NH_2$, $N C(O)CH_3$, $CH_2C(O)OH$,
25 SO_2NH_2 , $C(O)OH$, OCF_2H , isopropyl, C_2H_5OH , $C(O)OCH_3$, CH_2OH , NH-
 $CH=CH$, $HC=N-N-H$, $N=CH-S$, morpholinyl and $CH_2N(CH_3)_2$.

6. The compound of claim 3 wherein b is 0 and R² is H.
7. The compound of claim 4 wherein a is 1, R² is H and R is m-ethyl.
- 5 8. The compound of claim 4 wherein a is 1, and R is p-methoxy.
9. The compound of claim 4 wherein a is 1, and R is p-hydroxy.
- 10 10. The compound of claim 4 wherein a is 1, b is 0 and R is m-hydroxy.
11. The compound of claim 4 wherein a is 1, b is 0 and R is p-cyano.
12. The compound of claim 4 wherein a is 1, b is 0 and R is m-C(O)NH₂.
- 15 13. The compound of claim 4 wherein a is 1, b is 0 and R is p-NC(O)CH₃.
14. The compound of claim 4 wherein a is 1, b is 0 and R is p-CH₂C(O)OH.
- 20 15. The compound of claim 4 wherein a is 1, b is 0 and R is p-SO₂NH₂.
16. The compound of claim 4 wherein a is 1, b is 0 and R is p-CH₂OH.
17. The compound of claim 4 wherein a is 1, b is 0 and R is m-methoxy.
- 25 18. The compound of claim 4 wherein R is p-CH₂CH₂OH.
19. The compound of claim 4 wherein a is 1, b is 0 and R is HN-CH=CH.

20. The compound of claim 4 wherein a is 1, b is 0 and R is HC=N-NH.
21. The compound of claim 4 wherein a is 1, b is 0 and R is p-N-morpholinyl.
- 5 22. The compound of claim 4 wherein a is 1, b is 0 and R is N-CH-S.
23. The compound of claim 4 wherein a is 1, b is 0 and R is p-OCHF₂.
24. The compound of claim 4 wherein a is 2 and R is m-hydroxy and p-COOH.
- 10 25. The compound of claim 4 wherein a is 2 and R is m-hydroxy and p-CH₃.
26. The compound of claim 4 wherein a is 2 and R is m-hydroxy and p-OCH₃.
- 15 27. The compound of claim 4 wherein a is 2 and R is m-F and p-OCH₃.
28. The compound of claim 4 wherein b is 1 and R¹ is CH₃, a is 1 and R is m-hydroxy.
- 20 29. The compound of claim 4 wherein b is 1, R is CH₃, a is 1 and R is NH-N=CH.
30. The compound of claim 4 wherein b is 1, R is CH₃, a is 2 and R is N-CH=CH.
- 25 31. The compound of claim 4 wherein b is 1, R is CH₃, a is 1 and R is S-CH=N.
32. The compound of claim 4 wherein b is 1, R¹ is CH₃, a is 1 and R is p-hydroxy.

33. The compound of claim 4 wherein b is 1, R¹ is CH₃, a is 1 and R is p-CH₂OH.
- 5 34. The compound of claim 4 wherein b is 1, R¹ is CH₃, a is 2 and R is m-F and p-OCH₃.
35. The compound of claim 4 wherein b is 1, R¹ is Cl, a is 2 and R is m-hydroxy and p-OCH₃.
- 10 36. The compound of claim 4 wherein b is 1, R¹ is Cl, a is 2 and R is m, p-di-OCH₃.
37. The compound of claim 4 wherein b is 1, R¹ is Cl, a is 1 and R is HN-
15 N=CH.
38. The compound of claim 4 wherein b is 1, R¹ is Cl, a is 1 and R is p-N-morpholinyl.
- 20 39. The compound of claim 4 wherein b is 1, R¹ is Cl, a is 2 and R is m-CH₂N(C₂H₅)₂ and p-hydroxy.
40. The compound of claim 4 wherein b is 1, R¹ is Cl, a is 1 and R is p-CH₂OH.
- 25 41. A pharmaceutical composition comprising a pharmaceutically acceptable carrier or excipient and a compound according to Claim 1.
42. A compound represented by the general formula II:



wherein R^5 is selected from the group consisting of halogen, nitro, hydroxy, hydrocarbyl, substituted hydrocarbyl, amide, thioamide, amine, thioether and sulfonyl; R^3 is selected from the group consisting of D, halogen, nitro, hydroxy, hydrocarbyl, substituted hydrocarbyl, amide, thioamide, amine, thioether and sulfonyl and phosphonic acid; R^4 is selected from the group consisting of hydrogen, hydrocarbyl and substituted hydrocarbyl; b is 0 or an integer from 1 to 3; a is 0 or an integer of from 1 to 5; the wavy line represents a E or Z bond and pharmaceutically acceptable salts thereof provided however said compound is not

- 3-Phenylaminomethylene-1,3-dihydro-indol-2-one,
- 10 3-[(3-Bromo-phenylamino)-methylene]-1,3-dihydro-indol-2-one,
- 3-[(4-Bromo-phenylamino)-methyl]-1,3-dihydro-indol-2-one,
- 3-[(3-Bromo-phenylamino)-methyl]-1,3-dihydro-indol-2-one,
- 3-[(4-Ethyl-phenylamino)-methylene]-1,3-dihydro-indol-2-one,
- 3-Ethyl-phenylamino)-methylene]-1,3-dihydro-indol-2-one,
- 15 3-[(4-Methoxy-phenylamino)-methylene]-1,3-dihydro-indol-2-one,
- 4-[(2-Oxo-1,3-dihydro-indol-3-ylididenemethyl)-amino]-benzoic acid ethyl ester,
- 3-[(2-Ethyl-phenylamino)-methylene]-1,3-dihydro-indol-2-one,
- 3-[(3-Fluoro-4-methyl-phenylamino)-methylene]-1,3-dihydro-indol-2-one,

3-[(3-Fluoro-2-methyl-phenylamino)-methylene]-1,3-dihydro-indol-2-one,
3-[(4-Hydroxy-phenylamino)-methylene]-1,3-dihydro-indol-2-one,
3-[(3-Chloro-4-hydroxy-phenylamino)-methylene]-1,3-dihydro-indol-2-one or
3-[(4-Fluoro-2-methyl-phenylamino)-methylene]-1,3-dihydro-indol-2-one.

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43. A method for treating diseases related to unregulated tyrosine kinase signal transduction, wherein said disease is selected from the group consisting of carcinoma, sarcoma, leukemia, erythromblastoma, glioblastoma, meningioma, astrocytoma, melanoma, myoblastoma, brain cancer, bladder cancer, ovarian
10 cancer, gastric cancer, pancreas cancer, colon cancer, blood cancer, lung cancer, bone cancer, diabetic retinopathy, age-related macular degeneration, retinopathy of prematurity, arthritis and restenosis, hepatic cirrhosis, atherosclerosis and surgical adhesions. glomerulonephritis, diabetic nephropathy, malignant nephrosclerosis, thrombotic microangiopathy syndromes, transplant rejection and glomerulopathies,
15 psoriasis, diabetes mellitus, wound healing, inflammation and neurodegenerative diseases the method comprising the step of administering to a subject in need thereof a therapeutically effective amount of a compound of claim 1.

44. The method of claim 84 wherein said disease is diabetic retinopathy.

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45. The method of claim 84 wherein said disease is age-related macular degeneration.

46. The method of claim 84 wherein said disease is psoriasis.

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47. The compound of claim 1 wherein R is selected from the group consisting of fluoro, methyl, $\text{NR}_2(\text{CR}^7\text{R}^8)_d\text{R}^6$ and $(\text{CR}^7\text{R}^8)_c\text{R}^6$, wherein R^6 is selected from

- the group consisting of diloweralkylamino, piperidinyl, 3-fluoropyrrolidinyl, 3-fluoropiperidinyl, 2-pyridinyl, 3-pyridinyl, 4-pyridinyl, piperidinyl 3-pyrrolinyl, pyrrolidinyl, methyl isonipecotate, N-(2-methoxyethyl)-N-methylamyl, 1,2,3,6-tetrahydropyridinyl, morpholinyl, hexamethyleneiminyl, piperazinyl-2-one,
- 5 piperazinyl, N-(2-methoxyethyl)ethylaminyl, thiomorpholinyl, heptamethyleneiminyl, 1-piperazinylcarboxaldehyde, 2,3,6,7-tetrahydro-(1H)-1,4-diazepinyl-5(4H)-one, N-methylhomopiperazinyl, (dimethylamino)pyrrolidinyl, N-(2-methoxyethyl)-N-propylaminyl, Isoindolinyl, Nipecotamidinyl, isonipecotamidinyl, 1-acetylpiperazinyl, 3-acetamidopyrrolidinyl, trans-
- 10 decahydroisoquinolinyl, cis-decahydroisoquinolinyl, N-acetylhomopiperazinyl, 3-(diethylamino)pyrrolidinyl, 1,4-dioxo-8-azaspiro[4.5]decaninyl, 1-(2-methoxyethyl)-piperazinyl, 2-pyrrolidin-3-ylpyridinyl, 4-pyrrolidin-3-ylpyridinyl, 3-(methylsulfonyl)pyrrolidinyl, 3-picolylmethylaminyl, 2-(2-methylaminoethyl)pyridinyl, 1-(2-pyrimidyl)-piperazinyl,
- 15 1-(2-pyrazinyl)-piperazinyl, 2-methylaminomethyl-1,3-dioxolane, 2-(N-methyl-2-aminoethyl)-1,3-dioxolane, 3-(N-acetyl-N-methylamino)pyrrolidinyl, 2-methoxyethylaminyl, tetrahydrofurfurylaminyl, 4-aminotetrahydropyran, 2-amino-1-methoxybutane, 2-methoxyisopropylaminyl, 1-(3-aminopropyl)imidazole, histamyl, N,N-diisopropylethylenediaminyl, 1-benzyl-
- 20 3-aminopyrrolidyl 2-(aminomethyl)-5-methylpyrazinyl, 2,2-dimethyl-1,3-dioxolane-4-methanaminyl, (R)-3-amino-1-N-BOC-pyrrolidinyl, 4-amino-1,2,2,6,6-pentamethylpiperidinyl, 4-aminomethyltetrahydropyranyl, ethanolamine and alkyl-substituted derivatives thereof.
- 25 48. The compound of claim 47 wherein R⁶ is selected from the group consisting of dimethylamino, diethylamino, 3-fluoropyrrolidinyl, 3-fluoropiperidinyl, 3-pyridinyl, 4-pyridinyl, pyrrolidinyl, morpholinyl, piperazinyl, heptamethyleneiminyl, tetrahydrofurfurylaminyl, 4-aminotetrahydropyranyl, N,N-diisopropylethylenediaminyl and 4-aminomethyltetrahydropyranyl.

49. The compound of claim 48 wherein R⁶ is unsubstituted or mono or dimethyl substituted morpholinyl, pyrrolidinyl, piperazinyl or piperidinyl.
50. The compound of claim 49 wherein R⁶ is morpholinyl.
51. The method of claim 43 wherein R is selected from the group consisting of fluoro, methyl, NR₂(CR⁷R⁸)_dR⁶ and (CR⁷R⁸)_c-R⁶, wherein R⁶ is selected from the group consisting of diloweralkylamino, 3-fluoropyrrolidinyl, piperidinyl, 3-fluoropiperidinyl, 2-pyridinyl, 3-pyridinyl, 4-pyridinyl, 3-pyrrolinyl, pyrrolidinyl, methyl isonipecotate, N-(2-methoxyethyl)-N-methylamyl, 1,2,3,6-tetrahydropyridinyl, morpholinyl, hexamethyleneiminyl, piperazinyl-2-one, piperazinyl, N-(2-methoxyethyl)ethylaminyl, thiomorpholinyl, heptamethyleneiminyl, 1-piperazinylcarboxaldehyde, 2,3,6,7-tetrahydro-(1H)-1,4-diazepinyl-5(4H)-one, N-methylhomopiperazinyl, (dimethylamino)pyrrolidinyl, N-(2-methoxyethyl)-N-propylaminyl, isoindolinyl, nipecotamidinyl, isonipecotamidinyl, 1-acetylpiperazinyl, 3-acetamidopyrrolidinyl, trans-decahydroisoquinolinyl, cis-decahydroisoquinolinyl, N-acetylhomopiperazinyl, 3-(diethylamino)pyrrolidinyl, 1,4-dioxo-8-azaspiro[4.5]decanyl, 1-(2-methoxyethyl)-piperazinyl, 2-pyrrolidin-3-ylpyridinyl, 4-pyrrolidin-3-ylpyridinyl, 3-(methylsulfonyl)pyrrolidinyl, 3-picolylmethylaminyl, 2-(2-methylaminoethyl)pyridinyl, 1-(2-pyrimidyl)-piperazinyl, 1-(2-pyrazinyl)-piperazinyl, 2-methylaminomethyl-1,3-dioxolane, 2-(N-methyl-2-aminoethyl)-1,3-dioxolane, 3-(N-acetyl-N-methylamino)pyrrolidinyl, 2-methoxyethylaminyl, tetrahydrofurfurylaminyl, 4-aminotetrahydropyran, 2-amino-1-methoxybutane, 2-methoxyisopropylaminyl, 1-(3-aminopropyl)imidazole, histaminyl, N,N-diisopropylethylenediaminyl, 1-benzyl-3-aminopyrrolidyl 2-(aminomethyl)-5-methylpyrazinyl, 2,2-dimethyl-1,3-dioxolane-4-methanaminyl, (R)-3-amino-1-N-BOC-pyrrolidinyl, 4-amino-1,2,2,6,6-

pentamethylpiperidinyl, 4-aminomethyltetrahydropyranyl, ethanolamine and alkyl-substituted derivatives thereof.

52. The method of claim 51 wherein R⁶ is selected from the group consisting of
5 dimethylamino, diethylamino, 3-fluoropyrrolidinyl, 3-fluoropiperidinyl, 3-pyridinyl,
4-pyridinyl, pyrrolidinyl, morpholinyl, piperazinyl, heptamethyleneiminyl,
tetrahydrofurfurylaminy, 4-aminotetrahydropyranyl, N,N-
diisopropylethylenediaminyl and 4-aminomethyltetrahydropyranyl.
- 10 53. The method of claim 52 where R⁶ is unsubstituted or mono or di-methyl
substituted morpholinyl, pyrrolidinyl, piperazinyl or piperidinyl.
54. The method of claim 53 wherein R⁶ is morpholinyl.